# 80809 SEARCH REQUEST FORM

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## Scientific and Technical Information Center

Requester's Full Name: Lisa. V. Cook  Examiner #: 77/34 Date: 11/21/02  Art Unit: 1641 Phone Number 305-0808 Serial Number: 09/462-9-31  Mail Box and Bldg/Room Location: 09/462-9-31  Mail Box and Bldg/Room Location: 09/462-9-31  If more than one search is submitted, please prioritize searches in order of need.  **********************************									
Title of Invention; Bolated C	isteucalcin fra	agrunts							
Inventors (please provide full names): $\widehat{JL}$	LKa Hellman	η, ,	£						
Earliest Priority Filing Date:									
Shichire Search  Therference Search  Oligoner Search  Within Seg. Id.  Search positions  with gamma-corb  positions 17, 21, on  to bib Sheet  Amended Case  Point of Contact: Thomas G. Larson, Ph.D.  703-308-7309  CM1, Rm. 6 B 01	No. 2  b - 30  oxylated glub	int of Contact: Point of Contact: S G. Larson, Ph.D. 703-398-7309 CM1, Rm. 6 B 01	7-30 Claims						
STAFF USE ONLY  Searcher:	Type of Search  NA Sequence (#)  AA Sequence (#)  Structure (#)  Bibliographic  Litigation  Fulltext  Patent Family  Other	Vendors and cost where appl STN	G-Gened						

PTO-1590 (1-2000)

### Marked-Up Copy of the Amended Claims

25 (amended). A monoclonal antibody or recombinant antibody fragment [having the eapability of binding] which binds a human gamma-carboxylated osteocalcin fragment, wherein said monoclonal antibody or recombinant antibody fragment has the specificity to epitopes that have been identified on said gamma-carboxylated fragment of osteocalcin, said osteocalcin fragment selected from the group consisting of

- i) a fragment which spans from amino acid in position 7 to amino acid in position 30 of the amino acid sequence set forth in SEQ ID NO:2 in which all three glutamic acids in positions 17, 21 and 24 of said sequence are gamma-carboxylated, and
- ii) a fragment which spans from amino acid in position 6 to amino acid in position 30 of the amino acid sequence of SEQ ID NO:2[, and that] in which all three glutamic acids in the positions 17, 21 and 24 of said sequence are gamma-carboxylated.

26 (amended). A non-competitive immunoassay for quantitative determination of a gamma-carboxylated osteocalcin fragment in a sample comprising contacting [a] the sample [containing said osteocalcin fragment] with two monoclonal antibodies or recombinant antibody fragments which bind said osteocalcin fragment, [and detecting] measuring bound monoclonal antibody or recombinant antibody fragment, comparing said amount with an amount measured in a sample having a known quantity of said osteocalcin fragment, wherein said monoclonal antibody or recombinant antibody fragment has the specificity to epitopes that have been identified on said gamma-carboxylated fragment of osteocalcin, said osteocalcin fragment selected from the group consisting of

i) a fragment which spans from amino acid in position 7 to amino acid in position 30 of the amino acid sequence set forth in SEQ ID NO:2 in which all three glutamic acids in positions 17, 21 and 24 of said sequence are gamma-carboxylated, and

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ii) a fragment which spans from amino acid in position 6 to amino acid in position 30 of the amino acid sequence of SEQ ID NO:2[, and that] in which all three glutamic acids in the positions 17, 21 and 24 of said sequence are gamma-carboxylated.

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## Clean Copy of the Amended Claims

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5 (amended). A cell line producing the monoclonal antibody of claim 25.

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8 (twice amended). The immunoassay of claim 26, wherein the non-competitive immunoassay is carried out in either a one-step or a two-step incubation procedure.



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## \*BIBDATASHEET\*

Bib Data Sheet

**CONFIRMATION NO. 5666** 

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SERIAL NUMB 09/462,931		FILING DATE 01/18/2000 RULE	l	CLASS 435	GRC	DUP ART I 1641	UNIT	DO	ATTORNEY OCKET NO. 2328-115	
APPLICANTS								And the second s		
JUKKA HELLMA	N, TL	JRKU, FINLAND;								
SANNA-MARIA KAKONEN, TURKU, FINLAND; MATTI KARP, LITTOINEN, FINLAND;TIMO LOVGREN, TURKU, FINLAND; H. KALERVO VAANANEN, TURKU, FINLAND; KIM PETTERSSON, TURKU, FINLAND;										
** CONTINUING DATA **********************************										
** FOREIGN APPLICATIONS ************************************										
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ADDRESS 6449 ROTHWELL, FIGG, ERNST & MANBECK, P.C. 1425 K STREET, N.W. SUITE 800 WASHINGTON , DC 20005										
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Stic Search Subnitted 11/21/02. LYCOOK

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#### Marked-Up Copy of the Amended Claims

25 (amended). A monoclonal antibody or recombinant antibody fragment [having the capability of binding] which binds a human gamma-carboxylated osteocalcin fragment, wherein said monoclonal antibody or recombinant antibody fragment has the specificity to epitopes that have been identified on said gamma-carboxylated fragment of osteocalcin, said osteocalcin fragment selected from the group consisting of

- i) a fragment which spans from amino acid in position 7 to amino acid in position 30 of the amino acid sequence set forth in SEQ ID NO:2 in which all three glutamic acids in positions 17, 21 and 24 of said sequence are gamma-carboxylated, and
- ii) a fragment which spans from amino acid in position 6 to amino acid in position 30 of the amino acid sequence of SEQ ID NO:2[, and that] in which all three glutamic acids in the positions 17, 21 and 24 of said sequence are gamma-carboxylated.

26 (amended). A non-competitive immunoassay for quantitative determination of a gamma-carboxylated osteocalcin fragment in a sample comprising contacting [a] the sample [containing said osteocalcin fragment] with two monoclonal antibodies or recombinant antibody fragments which bind said osteocalcin fragment, [and detecting] measuring bound monoclonal antibody or recombinant antibody fragment, comparing said amount with an amount measured in a sample having a known quantity of said osteocalcin fragment, wherein said monoclonal antibody or recombinant antibody fragment has the specificity to epitopes that have been identified on said gamma-carboxylated fragment of osteocalcin, said osteocalcin fragment selected from the group consisting of

i) a fragment which spans from amino acid in position 7 to amino acid in position 30 of the amino acid sequence set forth in SEQ ID NO:2 in which all three glutamic acids in positions 17, 21 and 24 of said sequence are gamma-carboxylated, and

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ii) a fragment which spans from amino acid in position 6 to amino acid in position 30 of the amino acid sequence of SEQ ID NO:2[, and that] in which all three glutamic acids in the positions 17, 21 and 24 of said sequence are gamma-carboxylated.

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### **Clean Copy of Amended Claims**

25 (amended). A monoclonal antibody or recombinant antibody fragment which binds a human gamma-carboxylated osteocalcin fragment, said monoclonal antibody or recombinant antibody fragment has the specificity to epitopes that have been identified on said gamma-carboxylated fragment of osteocalcin, said osteocalcin fragment selected from the group consisting of:

- i) a fragment which comprises amino acids 7-30 of the amino acid sequence set forth in SEQ ID NO:2 in which all three glutamic acids in positions 17, 21, and 24 of said sequence are gamma-carboxylated, and
- ii) a fragment which comprises amino acids 6-30 of the amino acid sequence of SEQ ID NO:2 in which all three glutamic acids in the positions 17, 21 and 24 of said sequence gamma-carboxylated.

26 (amended). A non-competitive immunoassay for quantitative determination of a gamma-carboxylated osteocalcin fragment comprising contacting a sample containing said osteocalcin fragment with two monoclonal antibodies or recombinant antibody fragments which bind said osteocalcin fragment, detecting bound monoclonal antibody or recombinant antibody fragments and correlating bound monoclonal antibody or recombinant antibody fragments with amount of osteocalcin fragment present, wherein said monoclonal antibody or recombinant antibody fragment has the specificity to epitopes that have been identified on said gamma-carboxylated fragment of osteocalcin, said osteocalcin fragment selected from the group consisting of

i) a fragment which comprises amino acids 7-30 of the amino acid sequence set forth in SEQ ID NO:2 in which all three glutamic acids in positions 17, 21 and 24 of said sequence are gamma-carboxylated, and

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ii) a fragment which comprises amino acids 6-30 of the amino acid sequence of SEQ ID NO:2 in which all three glutamic acids in the positions 17, 21 and 24 of said sequence are gamma-carboxylated.

### **Clean Copy of Amended Claims**

25 (amended). A monoclonal antibody or recombinant antibody fragment which binds a human gamma-carboxylated osteocalcin fragment, said monoclonal antibody or recombinant antibody fragment has the specificity to epitopes that have been identified on said gamma-carboxylated fragment of osteocalcin, said osteocalcin fragment selected from the group consisting of:

- i) a fragment which comprises amino acids 7-30 of the amino acid sequence set forth in SEQ ID NO:2 in which all three glutamic acids in positions 17, 21, and 24 of said sequence are gamma-carboxylated, and
- ii) a fragment which comprises amino acids 6-30 of the amino acid sequence of SEQ ID NO:2 in which all three glutamic acids in the positions 17, 21 and 24 of said sequence gamma-carboxylated.

26 (amended). A non-competitive immunoassay for quantitative determination of a gamma-carboxylated osteocalcin fragment comprising contacting a sample containing said osteocalcin fragment with two monoclonal antibodies or recombinant antibody fragments which bind said osteocalcin fragment, detecting bound monoclonal antibody or recombinant antibody fragments and correlating bound monoclonal antibody or recombinant antibody fragments with amount of osteocalcin fragment present, wherein said monoclonal antibody or recombinant antibody fragment has the specificity to epitopes that have been identified on said gamma-carboxylated fragment of osteocalcin, said osteocalcin fragment selected from the group consisting of

i) a fragment which comprises amino acids 7-30 of the amino acid sequence set forth in SEQ ID NO:2 in which all three glutamic acids in positions 17, 21 and 24 of said sequence are gamma-carboxylated, and

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ii) a fragment which comprises amino acids 6-30 of the amino acid sequence of SEQ ID NO:2 in which all three glutamic acids in the positions 17, 21 and 24 of said sequence are gamma-carboxylated.

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## Clean Copy of the Amended Claims

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5 (amended). A cell line producing the monoclonal antibody of claim 25.

8 (twice amended). The immunoassay of claim 26, wherein the non-competitive immunoassay is carried out in either a one-step or a two-step incubation procedure.